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## FENT COOPERATION TREA

	From the INTERNATIONAL BUREAU
PCT	То:
NOTIFICATION OF ELECTION  (PCT Rule 61.2)	Assistant Commissioner for Patents United States Patent and Trademark Office Box PCT Washington, D.C.20231 ÉTATS-UNIS D'AMÉRIQUE
Date of mailing (day/month/year) 14 January 2000 (14.01.00)	in its capacity as elected Office
International application No. PCT/GB99/01557	Applicant's or agent's file reference PBA3/DO88155PWO
International filing date (day/month/year) 17 May 1999 (17.05.99)	Priority date (day/month/year) 16 May 1998 (16.05.98)
Applicant OLIVER, Stephen, George et al	
in a notice effecting later election filed with the Inte	ry Examining Authority on: 1999 (13.12.99)
The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland	Authorized officer  H. Zhou

Telephone No.: (41-22) 338.83.38

Facsimile No.: (41-22) 740.14.35

#### TENT COOPERATION TRE. Y

	From the	INTERNATIONAL BU	JREAU
PCT	To:		
NOTIFICATION OF THE RECORDING OF A CHANGE  (PCT Rule 92bis.1 and Administrative Instructions, Section 422)	Marks Susse: 83-85 Manch	ATKINSON, Peter, Birch Marks & Clerk Sussex House 83-85 Mosley Street Manchester M2 3LG ROYAUME-UNI	
09 October 2000 (09.10.00)			
Applicant's or agent's file reference PBA3/DO88155PWO		IMPORTANT NOTIF	FICATION
International application No. PCT/GB99/01557	1	al filing date (day/month/yea ay 1999 (17.05.99)	ar)
The following indications appeared on record concerning:      The applicant the inventor	the agent		n representative
Name and Address UNIVERSITY OF MANCHESTER Institute of Science and Technology		State of Nationality GB Telephone No.	State of Residence GB
P.O. Box 88 Manchester M60 1QD United Kingdom		Facsimile No.	·
	-	Teleprinter No.	
The International Bureau hereby notifies the applicant that the X the person the name the add		hange has been recorded c	concerning: the residence
Name and Address THE VICTORIA UNIVERSITY OF MANCHESTER		State of Nationality GB Telephone No.	State of Residence GB
Oxford Road Manchester M13 9PL United Kingdom		Facsimile No.	
		Teleprinter No.	
			<del></del>
3. Further observations, if necessary:			
4. A copy of this notification has been sent to:			
X the receiving Office	Г	the designated Offices of	concerned
the International Searching Authority	X	the elected Offices conc	erned
the International Preliminary Examining Authority		other:	
	Authorized of	fficer	
The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland		Sean Taylor	
Facsimile No.: (41-22) 740.14.35	Telephone N	o.: (41-22) 338.83.38	



### **PCT**

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#### INTERNATIONAL PRELIMINARY EXAMINATION REPORT

16

(PCT Article 36 and Rule 70)

	_		FOR FURTHER AC	See Notification of Transmittal of International  CTION Preliminary Examination Report (Form PCT/IPEA/416)			
PBA/DO	3815	5PWU				· · · · · · · · · · · · · · · · · · ·	
Internationa			International filing date (da	ay/month	/year)	Priority date (day/month/year)	
PCT/GB9	99/01	557	17/05/1999			16/05/1998	
Internationa C12N15/		ent Classification (IPC) or na	tional classification and IPC	•			
Applicant							
UNIVER	SITY	OF MANCHESTER IN	ISTITUTE OF et al.				
		ational preliminary exami smitted to the applicant a		repared	by this Inte	rnational Preliminary Examining Authority	
2. This f	REPC	ORT consists of a total of	8 sheets, including this	cover s	heet.		
This report is also accompanied by ANNEXES, i.e. sheets of the description, claims and/or drawings which hav been amended and are the basis for this report and/or sheets containing rectifications made before this Authority (see Rule 70.16 and Section 607 of the Administrative Instructions under the PCT). These annexes consist of a total of 3 sheets.							
3. This r	eport	contains indications rela	ting to the following item	s:			
1	$\boxtimes$	Basis of the report					
H	$\boxtimes$	Priority					
III		Non-establishment of o	pinion with regard to nov	elty, inv	entive step	and industrial applicability	
IV	$\boxtimes$	Lack of unity of invention	on				
V	×		nder Article 35(2) with reg ons suporting such stater		novelty, inve	entive step or industrial applicability;	
VI		Certain documents cité	ed				
VII		Certain defects in the in	nternational application				
VIII	Ø	Certain observations of	n the international applica	ation			
Date of sub	missio	on of the demand		Date of	completion of	this report	
			i			00	

Date of submission of the demand	Date of completion of this report	
13/12/1999	3 0. 0860	
Name and mailing address of the international preliminary examining authority:	Authorized officer	CRESISCHES MICHIGAN
European Patent Office D-80298 Munich Tel. +49 89 2399 - 0 Tx: 523656 epmu d	Marinoni, J-C	(in the same of th
Fax: +49 89 2399 - 4465	Telephone No. +49 89 2399 8563	distant. Min.

## INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/GB99/01557

I.	Bas	is o	f th	e re	port

••				
1.	resp	oonse to an invitatio	rawn on the basis of ( <i>substitute</i> on under Article 14 are referred o not contain amendments.):	sheets which have been furnished to the receiving Office in to in this report as "originally filed" and are not annexed to
	Des	cription, pages:		
	1-38	3	as originally filed	
	Clai	ims, No.:		
	1-2	1	as originally filed	
	22-3	38	with telefax of	11/08/2000
	Dra	wings, sheets:		
	1/9-	9/9	as originally filed	
2.	The	amendments have	e resulted in the cancellation of:	
		the description,	pages:	
		the claims,	Nos.:	
		the drawings,	sheets:	
3.			en established as if (some of) to beyond the disclosure as filed (f	ne amendments had not been made, since they have been Rule 70.2(c)):
4.	Ado	litional observation	s, if necessary:	
II.	Pric	ority		
1.		This report has be prescribed time lin		nad been claimed due to the failure to fumish within the
		☐ copy of the ea	arlier application whose priority	has been claimed.

 $\hfill\Box$  translation of the earlier application whose priority has been claimed.

## INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No. PCT/GB99/01557

2.		This report has been established as if no priority had been claimed due to the fact that the priority claim has been found invalid.
Th	us fo	or the purposes of this report, the international filing date indicated above is considered to be the relevant date
3.	Add	ditional observations, if necessary:
	see	separate sheet
		·
IV.	. Lac	ck of unity of invention
1.	ln r	esponse to the invitation to restrict or pay additional fees the applicant has:
		restricted the claims.
		paid additional fees.
		paid additional fees under protest.
		neither restricted nor paid additional fees.
2.	×	This Authority found that the requirement of unity of invention is not complied and chose, according to Rule 68.1, not to invite the applicant to restrict or pay additional fees.
3.	Thi	s Authority considers that the requirement of unity of invention in accordance with Rules 13.1, 13.2 and 13.3 is
		complied with.
	Ø	not complied with for the following reasons:
		see separate sheet
4.		nsequently, the following parts of the international application were the subject of international preliminary amination in establishing this report:
	×	all parts.
		the parts relating to claims Nos

#### INTERNATIONAL PRELIMINARY **EXAMINATION REPORT**

International application No. PCT/GB99/01557

V. Reasoned stat m nt under Articl 35(2) with regard to nov lty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Statement

Novelty (N)

Yes:

Claims 1-38

Claims NONE No:

Inventive step (IS)

Yes:

Claims 1-35, 37, 38

No:

Claims 36

Industrial applicability (IA)

Yes:

Claims 1-38

No:

Claims NONE

2. Citations and explanations

see separate sheet

#### VIII. Certain observations on the international application

The following observations on the clarity of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made:

see separate sheet

#### Re Item II

#### **Priority**

The document ZHANG et al. 'Down-regulation of the expression of PKC1 and SRB1/PSA1/VIG9, two genes involved in cell wall integrity in *Saccharomyces cerevisiae*, causes flocculation.' MICROBIOLOGY, Vol. 145, No. Pt 2, February 1999, pages 309-316, has been cited as a P document.

The International Preliminary Examination Authority considers the priority of the present application to be valid. Therefore, this document is not taken into consideration for the establishement of the following opinion. However, some of the documents therein cited are considered to be relevant and are fully citable.

#### Re Item IV

#### Lack of unity of invention

Reference is made to the following document:

D1: J. CELL BIOL., Vol. 116, No. 5, 1992, pages 1221-1229, Levin and Bartlett-Heubusch 'Mutants in the S. cerevisiae PKC1 gene display a cell specific osmotic stability defect'

Reference is made to the following document which was not cited in the search report (cited in Zhang et al. 1999).

- **D2**: Zhang, N., PhD thesis, University of Manchester-Institute of Science and Technology, 1997.
- 1. The separate inventions/groups of invention are:
  - Yeast cells wherein the SRB1 and PKC1 genes are each under the control of a heterologous (inducible) promoter, method of regulating yeast cell lysis and method of fermentation using said yeast cells (claims 1-14, 32-34, 37, 38),
  - (ii) Yeast cells wherein the PKC1 gene is under the control of a heterologous inducible promoter, method of regulating yeast cell flocculation using said yeast cells (claims 15-23, 35)
  - (iii) Yeast cells wherein the SRB1 gene is under the control of a heterologous promoter, method of fermentation using said yeast cells (claims 24-31, 36)

- They are not so linked as to form a single general inventive concept (Rule 13.1 2. PCT) for the following reasons:
  - (a) the technical feature linking group (i) and (ii) of inventions resides in yeast cells comprising the PKC1 gene under the control of an inducible heterologous promoter; this technical feature is not novel (see D1).
  - the technical feature linking group (i) and (iii) of inventions resides in yeast cells comprising the SRB1 gene under the control of a heterologous promoter (i.e. subject-matter of claim 36); this subject-matter is not novel (see the grounds for this objection)
  - the technical feature linking group (ii) and (iii) of inventions resides in yeast cells; this feature is obviously not new.

#### Re Item V

Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

- Claim 1 is directed to a yeast cell containing the SRB1/PSA1 gene and the PKC1 1. gene or functional derivatives thereof each operatively linked to a heterologous inducible promoter.
  - None of the available document discloses nor suggests such cells or methods using said cells. Therefore, the subject-matter of claim 1 meets the requirements of Article 33(2) PCT concerning novelty and Article 33(3) PCT concerning inventive step.
  - Consequently, claims 2-14 and 32-34 also meet these requirements of the PCT.
- Claim 15 is directed to a method of regulating yeast cell flocculation comprising 2. growing yeast cells containing the PKC1 gene operatively linked to an inducible promoter. None of the available document discloses nor suggests a link between PKC1 and yeast flocculation.
  - None of the available documents neither discloses nor suggests such a method. Therefore, the subject-matter of claim 15 meets the requirement of Article 33(2) PCT concerning novelty and of Article 33(3) PCT concerning inventive step.

Consequently, dependent claims 16-23 also meet these requirements of the PCT.

Claim 24 is directed to a method of fermentation comprising growing yeast cells 3. containing the SRB1/PSA1 gene under the control of a heterologous promoter. None of the available documents neither discloses nor suggests such a method. Therefore, claim 24 meets the requirement of Article 33(3) PCT concerning inventive step.

Consequently, dependent claims 25-31 also meet these requirements of the PCT.

Claim 35 is directed to yeast cells comprising the PKC1 gene under the control of 4. an inducible heterologous promoter and selected from two cell lines transformed with specific constructs, and a third cell line. None of the available documents suggests that the specific cells lines containing one of the specific constructs could or should be obtained.

Therefore, the subject-matter of claim 35 meets the requirements of Article 33(3) PCT concerning inventive step.

Claim 36 is directed to yeast cells comprising the SRB1 gene under the control of 5. a heterologous promoter.

Zhang et al. (1999) refers (see page 310, lines 45-55) to document D2 which appears to disclose yeast cells transformed with vectors containing the SRB1/PSA1 gene under the control of the MET3 promoter.

Therefore, the subject-matter of claim 36 does not meet the requirements of Article 33(2) PCT concerning novelty.

#### Re Item VIII

#### Certain observations on the international application

- Many claims refer to names of plasmids or cells lines which appear to be arbitrary 1. denomination which have no defined meaning for the skilled person. Said claim: (see claims 6-8, 20-22, 29-31, 35) are therefore not clear (Article 6 PCT).
- Many claims refer to "functional derivatives" of the SRB1 and/or PKC1 genes. 2. None of these functional derivatives is examplified in the application as filed. The

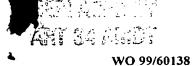


### INTERNATIONAL PRELIMINARY

International application No. PCT/GB99/01557

**EXAMINATION REPORT - SEPARATE SHEET** 

scope of protection afforded by the term "functional derivative" is not fully clear (Article 6 PCT). Furthermore, none of the examples in the application as filed refers specifically to one of such "functional derivatives" (Article 6 PCT). It appears that the caracterisation of such "functional derivatives" useful in the claimed processes would place undue burden on the skilled person (Article 5 PCT).



22. The method according to any one of claims 15 to 20 wherein the yeast cells are ZO123 or ZO124 transformed with the *PKC1* gene or functional derivative thereof operatively linked to an inducable promoter.

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- 23. The method according to any one of claims 15 to 22 for increasing the sedimentation of yeast cells or cell ghosts / debris form a medium within which the yeast cells are grown.
- 23. A method of fermentation comprising growing yeast cells containing the *SRB1/PSA1* gene or functional derivative thereof operatively linked to a heterologous promoter in a growth medium in which *SRB1/PSA1* expression is regulated by the heterologous promoter whereby said cells flocculate.
- 24. The method according to claim 23 wherein the yeast cell is a strain of Saccharomyces cerevisiae
- 25. The method according to claim 23 wherein the yeast cell is a strain of *Pichia pastoris*. Hansenula polymorpha or Kluyveromyces lactis.
- 26. The method according to any one of claims 23 25 wherein the *SRB1/PSA1* gene or functional derivative thereof is operatively linked to a methionine regulated promoter.
- 27. The method according to claim 26 wherein the methionine regulated promoter is pMET3.
- 28. The method according to claim 27 wherein the *SRB1/PSA1* gene or functional derivatives thereof operatively linked to an inducable promoter is derived from the recombinant vector SRB1.9e.

- 29. The method according to claim 28 wherein the yeast cells are ZO-125.
- 30. The method according to claim 28 wherein the yeast cells are FY23SRB1MET3.
- A method of fermentation comprising growing yeast cells containing the *SRB1/PSA1* and *PKC1* gene or functional derivatives thereof operatively linked to a heterologous promoter in a growth medium in which *SRB1/PSA1* and *PKC1* expression is regulated by the heterologous promoter whereby said cells flocculate.
- 32. The method according to claim 31 wherein the cells are cells according to any one of claims 1 8.
- 33. The method according to claim 31 wherein the cells contain the *PKC1* gene or a functional derivative thereof operatively linked to a heterologous inducable promoter and the *SRB1/PSA1* gene or a functional derivative thereof operatively linked to a heterologous promoter.
- 34. A yeast cell containing the *PKC1* gene or functional derivatives thereof operatively linked to a heterologous inducable promoter.
- 35. A yeast cell containing the *SRB1/PSA1* gene or functional derivatives thereof operatively linked to a heterologous promoter.
- 36. A yeast cell containing the *PKC1* gene or a functional derivative thereof operatively linked to a heterologous inducable promoter and the *SRB1/PSA1* gene or a functional derivative thereof operatively linked to a heterologous promoter.
- 37. A yeast cell according to any one of claims 34 36 wherein the promoter or promoters is/are pMET3.

#### INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference	FOR FURTHER see Notification of Transmittal of International Search Report (Form PCT/ISA/220) as well as, where applicable, item 5 below.					
PBA3/D088155PW0	ACTION					
International application No.	International filing date (day/month/year) (Earliest) Priority Date (day/month/year)					
PCT/GB 99/01557	17/05/1999 16/05/1998					
Applicant						
UNIVERSITY OF MANCHESTER	et al.					
This International Search Report has been according to Article 18. A copy is being tra	n prepared by this International Searching Auth Insmitted to the International Bureau.	ority and is transmitted to the applicant				
This International Search Report consists	of a total of5sheets.					
	a copy of each prior art document cited in this	report.				
1. Basis of the report						
	international search was carried out on the bas ess otherwise indicated under this item.	is of the international application in the				
the international search w Authority (Rule 23.1(b)).	as carried out on the basis of a translation of the	ne international application furnished to this				
		ternational application, the international search				
was carried out on the basis of the X contained in the internatio	e sequence listing : nal application in written form.					
	rnational application in computer readable form	1.				
furnished subsequently to	this Authority in written form.					
furnished subsequently to	furnished subsequently to this Authority in computer readble form.					
X the statement that the info	rmation recorded in computer readable form is	identical to the written sequence listing has been				
2. Certain claims were four	nd unsearchable (See Box I).					
3. Unity of invention is lack	king (see Box II).					
4. With regard to the title,						
``	the text is approved as submitted by the applicant.					
	[X] the text has been established by this Authority to read as follows:  REGULATED EXPRESSION OF PKC AND/OR SRB1/PSA1 IN YEAST					
REGOLATED EXTRESSION C	I INC AND ON SUBILIBATING	LNJI				
5. With regard to the abstract,	•					
X the text is approved as sul	bmitted by the applicant.					
	ned, according to Rule 38.2(b), by this Authorit date of mailing of this international search rep					
6. The figure of the drawings to be publi		- authority.				
		None of the figures				
	as suggested by the applicant.  None of the figures.					
	because the applicant failed to suggest a figure.  because this figure better characterizes the invention.					

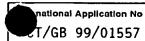
#### INTERNATIONAL SEARCH REPORT



a. classi IPC 6	FICATION OF SUBJECT MATTER C12N15/81 C12N15/54 C12N	11/19	//(C12N1/19,C12R1	:865)
According to	o International Patent Classification (IPC) or to both national c	lassification ar	d IPC	
	SEARCHED			
Minimum do	cumentation searched (classification system followed by class C12N C12R	ssification sym	ools)	
Documenta	ion searched other than minimum documentation to the exter	nt that such do	cuments are included in the fields se	earched
Electronic d	ata base consulted during the international search (name of o	data base and,	where practical, search terms used	
C. DOCUM	ENTS CONSIDERED TO BE RELEVANT			
Category °	Citation of document, with indication, where appropriate, of	the relevant p	assages	Relevant to claim No.
P,X	ZHANG N. ET AL.: "Down-regul expression of PKC1 and SRB1/P genes involved in cell wall i Saccharomyces cerevisiae, cau flocculation." MICROBIOLOGY, vol. 145, no. Pt 2, February 1999 (1999-02), page XP000052921 / the whole document	SA1/VIGS ntegrity ses	9, two / in	1-37
X Furth	ner documents are listed in the continuation of box C.	X	Patent family members are listed	in annex.
"A" docume consid "E" earlier of filling d "L" docume which citation "O" docume other r "P" docume later th	nt which may throw doubts on priority claim(s) or is cited to establish the publication date of another in or other special reason (as specified) and referring to an oral disclosure, use, exhibition or	o oi in "X" do c: in "Y" do c d d m m in	or document published after the interpriority date and not in conflict with ted to understand the principle or the vention current of particular relevance; the cannot be considered novel or cannot volve an inventive step when the document of particular relevance; the cannot be considered to involve an inventive step when the document is combined with one or molents, such combined with one or molents, such combination being obvious the art.  current member of the same patent of the official particular search of the international search of the international search of the same patent of the same pa	the application but sory underlying the laimed invention be considered to current is taken alone laimed invention rentive step when the re other such docu-us to a person skilled family
Name and n	nailing address of the ISA  European Patent Office, P.B. 5818 Patentlaan 2  NL – 2280 HV Rijswijk  Tel. (+31-70) 340–2040, Tx. 31 651 epo nl,	A	thorized officer	

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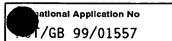
#### **INTERNATIONAL SEARCH REPORT**



	C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT					
Category °	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.			
Υ	BELDA F. & ZARATE V.: "Isolation and characterization of Schizosaccharomyces pombe fragile mutants" YEAST,		23			
	vol. 12, 1996, pages 555-564, XP000852922 <b>*</b> the whole document	:	1.00			
A			1-22, 24-37			
<b>Y</b>	WO 94 03609 A (IMP CANCER RES TECH ;PARKER PETER JOSEPH JACQUES (GB); GOODE NIGEL) 17 February 1994 (1994-02-17) abstract	·	23			
	page 16 -page 18 claims 1-4					
A			1-22, 24-37			
Α	WO 92 01798 A (UNIV MANCHESTER)/ 6 February 1992 (1992-02-06) abstract		1-37			
A	RECH S.B. ET AL.: "Complementation of the Saccharomyces cerevisiae srb1-1 mutation: an autoselection system for stable plasmid maintenance" CURR. GENET., vol. 21, 1992, pages 339-344, XP002122385	·	1-37			
	the whole document					
A	STATEVA L.I. ET AL.: "Cloning and characterization of a gene which determines osmotic stability in Saccharomyces cerevisiae" MOL. CELL. BIOL.,	,	1-37			
	vol. 11, no. 8, August 1991 (1991-08), pages 4235-4243, XP002122386 / the whole document	*				
<b>A</b>	CARO L.H.P. ET AL.: "Transcription of multiple cell wall protein-encoding genes in Saccahromyces cerevisiae is differentially regulated during the cell cycle" FEMS MICROBIOL. LETTERS,		1-37			
·	vol. 161, no. 2, 15 April 1998 (1998-04-15), pages 345-349, XP000852924 / the whole document					
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#### INTERNATIONAL SEARCH REPORT



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C.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Α	STATEVA L.I. ET AL.: "Polyploid fragile strains of Saccharomyces cerevisiae a novel source of proteins for nutritional purposes" YEAST, vol. 4, no. 3, September 1988 (1988-09), pages 219-225, XP000852925  the whole document	1-37
<b>A</b>	VERNA J.: "A family of genes required for maintenance of cell wall integrity and for the stress response in Saccharomyces cerevisiae" PROC. NATL. ACAD. SCI. USA, vol. 94, December 1997 (1997-12), pages 13804-13809, XP002122387 for the whole document	1-37
<b>A</b>	KLIS F M: "REVIEW: CELL WALL ASSEMBLY IN YEAST" YEAST,GB,CHICHESTER, SUSSEX, vol. 10, page 851-869 XP000196331 ISSN: 0749-503X	1-37
<b>A</b>	WO 96 02629 A (UNIV MADRID COMPLUTENSE ( ;NOMBELA CANO CESAR (ES); ALVAREZ ALVAREZ) 1 February 1996 (1996-02-01) abstract	1-37
A .	SHANKAR C.S. ET AL.: "MIG1 overexpression causes flocculation in Saccharomyces cerevisiae" MICROBIOLOGY, vol. 142, no. Pt 9, 1996, pages 2663-2667, XP002122388/ the whole document	1-37
Α	US 5 716 808 A (RAYMOND CHRISTOPHER K) 10 February 1998 (1998-02-10) abstract	1-37
A	WO 88 10308 A (WHITEHEAD BIOMEDICAL INST) 29 December 1988 (1988-12-29) abstract/	1-37
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#### INTERNATIONAL SEARCH REPORT

ational Application No /GB 99/01557

C.(Continua	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		<del></del>	
Category °	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.	
A	DATABASE GENBANK 'Online! Accession No. X06413, 12 September 1993 (1993-09-12) CHEREST H. ET AL.: "Yeast MET3 gene for ATP sulphurylase" XP002122389/ cited in the application the whole document & CHEREST H. ET AL.: "The Saccharomyces cerevisiae MET3 gene: nucleotide sequence and relationship of the 5' non-coding region to that of MET25." MOL. GEN. GENET., vol. 210, no. 2, 1987, pages 307-313,	1-37		
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#### INTERNATIONAL SEARCH REPORT

tion on patent family members

national Application No I/GB 99/01557

Patent document cited in search report		Publication date	Patent family member(s)		Publication date
WO 9403609	Α	17-02-1994	EP JP	0658202 A 8503124 T	21-06-1995 09-04-1996
WO 9201798	Α	06-02-1992	AU	8296291 A	18-02-1992
WO 9602629	A	01-02-1996	ES AU	2092439 A 3078795 A	16-11-1996 16-02-1996
US 5716808	A	10-02-1998	AU CA CA EP EP WO US EP US	1158297 A 7673796 A 2237039 A 2237120 A 0889966 A 0862640 A 9717450 A 9717451 A 5965389 A 0920525 A 5854039 A	29-05-1997 29-05-1997 15-05-1997 15-05-1997 13-01-1999 09-09-1998 15-05-1997 15-05-1997 12-10-1999 09-06-1999 29-12-1998
W0 8810308	A	29-12-1988	US	5063154 A	05-11-1991

#### PATENT COOPERATION TREATY

From the INTERNATIONAL SEARCHING AUTHORITY	PCT						
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Applicant's or agent's file reference PBA3/D088155PW0	REPLY DUE  See paragraph 1 below						
International application No. PCT/GB 99/01557	International filing date (day/month/year) 17/05/1999						
Applicant UNIVERSITY OF MANCHESTER et al.	•						
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1. REPLY DUE within	days from the above date of maining						
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With regard to your Application PCT/6 for which we have received your Fax of dispatch of Cited document XPO0005292	BB99/01557 (Your Ref:PBA/PCB/JK/D88155PWO) of 3 December 1999 and concerning the 21 in error in place of XP000852921.						
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We apologise for any inconvienience of	caused.						
Yours sincerely,							

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Authorized officer

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